(12) INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

(19) World Intellectual Property Organization International Bureau

MIPO OMPL



(43) International Publication Date 17 June 2004 (17.06.2004)

PCT

(10) International Publication Number WO 2004/050867 A1

- (51) International Patent Classification⁷: C12N 15/00, 15/63, 15/74, C07H 21/00, 21/04, C07K 16/00
- (21) International Application Number:

PCT/US2002/038414

- (22) International Filing Date: 2 December 2002 (02.12.2002)
- (25) Filing Language:

English

(26) Publication Language:

English

- (71) Applicant (for all designated States except US): SEAT-TLE GENETICS, INC. [US/US]; 21823 30th Drive, S.E., Bothell, WA 98021 (US).
- (72) Inventors; and
- (75) Inventors/Applicants (for US only): MCDONAGH,

Charlotte, F. [GB/US]; 8018 28th Avenue NW, Seattle, WA 98117 (US). FRANCISCO, Joseph, A. [US/US]; 21705 92nd Avenue West, Edmonds, WA 98020 (US).

- (74) Agents: ANTLER, Adrian, M. et al.; Pennie & Edmonds LLP, 1155 Avenue of the Americas, New York, NY 10036 (US).
- (81) Designated States (national): CA, US.

Published:

with international search report

For two-letter codes and other abbreviations, refer to the "Guidance Notes on Codes and Abbreviations" appearing at the beginning of each regular issue of the PCT Gazette.

(54) Title: MODIFIED L49-sFv EXHIBITING INCREASED STABILITY AND METHODS OF USE THEREOF

(57) Abstract: The present invention relates to a modified L49 single chain antibody (L49-sFv) that exhibits increased refolding efficiency and/or greater stability in mouse serum, and surprisingly substantially maintains binding affinity for its binding ligand, p97 melanotransferrin. p97 melanotransferrin is expressed on the surface of a number of types of cancer (carcinoma) cells, e.g., melanoma cells, lung cancer cells, renal cancer cells, colon cancer cells. The present invention also relates to a modified L49-sFv fused or conjugated to a therapeutic agent, such as a cytotoxic molecule or a pro-drug converting enzyme. The present invention also relates to methods of using the modified L49-sFv molecules fused or conjugated to a therapeutic agent for treatment and/or prophylaxis of cancer, which cancer cells express p97 melanotransferrin.